

# Synthesis of new p-tert-butylcalix[4]arene-based polyammonium triazolyl amphiphiles and their binding with nucleoside phosphates

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## Abstract

© 2018 Burilov et al. The synthesis of new calix[4]arenes adopting a cone stereoisomeric form bearing two or four azide fragments on the upper rim and water-soluble triazolyl amphiphilic receptors with two or four polyammonium headgroups via copper-catalyzed azide-alkyne cycloaddition reaction has been performed for the first time. It was found that the synthesized macrocycles form stable aggregates with hydrodynamic diameters between 150–200 nm and electrokinetic potentials about +40 to +60 mV in water solutions. Critical aggregation concentration (CAC) values were measured using a micelle method with pyrene and eosin Y as dye probes. The CAC values of tetraalkyl-substituted macrocycles 12a,b (5  $\mu$ M for both) are significantly lower than those for dialkyl-substituted macrocycles 10a,b (790 and 160  $\mu$ M, respectively). Premicellar aggregates of macrocycles 10a,b and 12a,b with the dye eosin Y were used for nucleotides sensing through a dye replacement procedure. It is unusual that disubstituted macrocycles 10a,b bind more effectively a less charged adenosine 5'-diphosphate (ADP) than adenosine 5'-triphosphate (ATP). A simple colorimetric method based on polydiacetylene vesicles decorated with 10b was elaborated for the naked-eye detection of ADP with a detection limit of 0.5 mM.

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## Keywords

ADP, Amphiphile, ATP, Calix[4]arene, CuAAC, Eosin Y probe, Molecular recognition, Polydiacetylene, Self-assembly, Triazole

## References

- [1] Ngo, H. T.; Liu, X.; Jolliffe, K. A. *Chem. Soc. Rev.* 2012, 41, 4928–4965. doi:10.1039/C2CS35087D
- [2] García-España, E.; Díaz, P.; Llinares, J. M.; Bianchi, A. *Coord. Chem. Rev.* 2006, 250, 2952–2986. doi:10.1016/j.ccr.2006.05.018
- [3] Li, A.-F.; Wang, J.-H.; Wang, F.; Jiang, Y.-B. *Chem. Soc. Rev.* 2010, 39, 3729–3745. doi:10.1039/b926160p
- [4] Zhou, Y.; Xu, Z.; Yoon, Y. *Chem. Soc. Rev.* 2011, 40, 2222–2235. doi:10.1039/c0cs00169d
- [5] Ng, S.; Lim, H. S.; Ma, Q.; Gao, Z. *Theranostics* 2016, 6, 1683–1702. doi:10.7150/thno.15850
- [6] Liemburg-Apers, D. C.; Imamura, H.; Forkink, M.; Nooteboom, M.; Swarts, H. G.; Brock, R.; Smeitink, J. A. M.; Willems, P. H. G. M.; Koopman, W. J. H. *Pharm. Res.* 2011, 28, 2745–2757. doi:10.1007/s11095-011-0492-8

- [7] Molz, S.; Tharine, D.-C.; Decker, H.; Tasca, C. I. *Brain Res.* 2008, 1231, 113–120. doi:10.1016/j.brainres.2008.07.009
- [8] Perrault, D. M.; Chen, X.; Anslyn, E. V. *Tetrahedron* 1995, 51, 353–362. doi:10.1016/0040-4020(94)00901-6
- [9] Bazzicalupi, C.; Bencini, A.; Berni, E.; Bianchi, A.; Fornasari, P.; Giorgi, C.; Masotti, A.; Paoletti, P.; Valtancoli, B. *J. Phys. Org. Chem.* 2001, 14, 432–443. doi:10.1002/poc.385
- [10] Nakai, C.; Glinsmann, W. *Biochemistry* 1977, 16, 5636–5641. doi:10.1021/bi00644a039
- [11] Kuchelmeister, H. Y.; Schmuck, C. *Chem. – Eur. J.* 2011, 17, 5311–5318. doi:10.1002/chem.201003393
- [12] Mascaros, P. A.; Bazzicalupi, C.; Bianchi, A.; Giorgi, C.; Valero, M. D. G.; Garzón, R. L.; Salido, M. L. G.; Valtancoli, B. *Chem. Commun.* 2011, 47, 2814–2816. doi:10.1039/C0CC05054G
- [13] Sansone, F.; Baldini, L.; Casnati, A.; Ungaro, R. *New J. Chem.* 2010, 34, 2715–2728. doi:10.1039/c0nj00285b
- [14] Wenz, G.; Han, B.-H.; Müller, A. *Chem. Rev.* 2006, 106, 782–817. doi:10.1021/cr970027+
- [15] Lagona, J.; Mukhopadhyay, P.; Chakrabarti, S.; Isaacs, L. *Angew. Chem., Int. Ed.* 2005, 44, 4844–4870. doi:10.1002/anie.200460675
- [16] Yakimova, L. S.; Shurpik, D. N.; Stoikov, I. I. *Chem. Commun.* 2016, 52, 12462–12465. doi:10.1039/c6cc05797g
- [17] Antipin, I. S.; Kazakova, E. K.; Habicher, W. D.; Konovalov, A. I. *Russ. Chem. Rev.* 1998, 67, 905–922. doi:10.1070/RC1998v067n11ABEH000472
- [18] Stoikov, I. I.; Repejko, S. A.; Antipin, I. S.; Konovalov, A. I. *Heteroat. Chem.* 2000, 11, 518–527. doi:10.1002/1098-1071(2000)11:7<518::AID-HC10>3.0.CO;2-#
- [19] Neri, P.; Sessler, J. L.; Wang, M.-X. *Calixarenes and Beyond*; Springer: Netherlands, 2016; p 1062.
- [20] Vavilova, A. A.; Stoikov, I. I. *Beilstein J. Org. Chem.* 2017, 13, 1940–1949. doi:10.3762/bjoc.13.188
- [21] Solovieva, S. E.; Burilov, V. A.; Antipin, I. S. *Macroheterocycles* 2017, 10, 134–146. doi:10.6060/mhc170512a
- [22] Burilov, V.; Valiyakhmetova, A.; Mironova, D.; Sultanova, E.; Evtugyn, V.; Osin, Y.; Katsyuba, S.; Burganov, T.; Solovieva, S.; Antipin, I. *New J. Chem.* 2018, 42, 2942–2951. doi:10.1039/c7nj04099g
- [23] Shinkai, S.; Mori, S.; Koreishi, H.; Tsubaki, T.; Manabe, O. *J. Am. Chem. Soc.* 1986, 108, 2409–2416. doi:10.1021/ja00269a045
- [24] Zadmard, R.; Schrader, T. *J. Am. Chem. Soc.* 2005, 127, 904–915. doi:10.1021/ja045785d
- [25] Ibragimova, R. R.; Burilov, V. A.; Aimetdinov, A. R.; Mironova, D. A.; Evtugyn, V. G.; Osin, Y. N.; Solovieva, S. E.; Antipin, I. S. *Macroheterocycles* 2016, 9, 433–441. doi:10.6060/mhc161180b
- [26] Burilov, V. A.; Mironova, D. A.; Ibragimova, R. R.; Evtugyn, V. G.; Osin, Y. N.; Solovieva, S. E.; Antipin, I. S. *Bionanosci.* 2018, 8, 337–343. doi:10.1007/s12668-017-0484-1
- [27] Song, M.; Sun, Z.; Han, C.; Tian, D.; Li, H.; Kim, J. S. *Chem. – Asian J.* 2014, 9, 2344–2357. doi:10.1002/asia.201400024
- [28] Rostovtsev, V. V.; Green, L. G.; Fokin, V. V.; Sharpless, K. B. *Angew. Chem., Int. Ed.* 2002, 41, 2596–2599. doi:10.1002/1521-3773(20020715)41:14<2596::AID-ANIE2596>3.0.CO;2-4
- [29] Smith, P. A. S. *The Chemistry of Open-chain Organic Nitrogen Compounds*, Band 2, XII; Verlag W. A. Benjamin Inc.: New York-Amsterdam, 1966; p 531.
- [30] Fujii, S.; Nishina, K.; Yamada, S.; Mochizuki, S.; Ohta, N.; Atsushi, T.; Sakurai, K. *Soft Matter* 2014, 10, 8216–8223. doi:10.1039/c4sm01355g
- [31] Bew, S. P.; Brimage, R. A.; L'Hermit, N.; Sharma, S. V. *Org. Lett.* 2007, 9, 3713–3716. doi:10.1021/ol071047t
- [32] Burilov, V. A.; Nugmanov, R. I.; Ibragimova, R. R.; Solovieva, S. E.; Antipin, I. S.; Konovalov, A. I. *Mendeleev Commun.* 2013, 23, 113–115. doi:10.1016/j.mencom.2013.03.022
- [33] Verboom, W.; Durie, A.; Egberink, R. J. M.; Asfari, Z.; Reinhoudt, D. N. *J. Org. Chem.* 1992, 57, 1313–1316. doi:10.1021/jo00030a050
- [34] Kenis, P. J. A.; Noordman, O. F. J.; Schönherr, H.; Kerver, E. G.; Snellink-Ruël, B. H. M.; van Hummel, G. J.; Harkema, S.; van der Vorst, C. P. J. M.; Hare, J.; Picken, S. J.; Engbersen, J. F. J.; van Hulst, N. F.; Vancso, G. J.; Reinhoudt, D. N. *Chem. – Eur. J.* 1998, 4, 1125–1134. doi:10.1002/(SICI)1521-3765(19980710)4:7<1225::AL-CHEM1225>3.0.CO;2-6
- [35] Dordea, C.; Brisach, F.; Haddaoui, J.; Arnaud-Neu, F.; Bolte, M.; Casnati, A.; Böhmer, V. *Supramol. Chem.* 2010, 22, 347–357. doi:10.1080/10610271003678511
- [36] Tomapatanaget, B.; Tuntulani, T. *Tetrahedron Lett.* 2001, 42, 8105–8109. doi:10.1016/S0040-4039(01)01722-1
- [37] Maurin, A.; Varatharajan, S.; Colasson, B.; Reinaud, O. *Org. Lett.* 2014, 16, 5426–5429. doi:10.1021/ol502650c
- [38] Buttress, J. P.; Day, D. P.; Courtney, J. M.; Lawrence, E. J.; Hughes, D. L.; Blagg, R. J.; Crossley, A.; Matthews, S. E.; Redshaw, C.; Page, P. C. B.; Wildgoose, G. G. *Langmuir* 2016, 32, 7806–7813. doi:10.1021/acs.langmuir.6b02222
- [39] Li, Y.; Hoskins, J. N.; Sreerama, S. G.; Grayson, S. M. *Macromolecules* 2010, 43, 6225–6228. doi:10.1021/ma100599n

- [40] Aguiar, J.; Carpana, P.; Molina-Bolíver, J. A.; Carnero Ruiz, C. J. *Colloid Interface Sci.* 2003, 258, 116–122. doi:10.1016/S0021-9797(02)00082-6
- [41] Hunter, R. J. *Foundations of Colloid Science*; Oxford University Press: Oxford, 1987; p 673.
- [42] Patist, A.; Bhagwat, S. S.; Penfield, K. W.; Aikens, P.; Shah, D. O. J. *Surfactants Deterg.* 2000, 3, 53–58. doi:10.1007/s11743-000-0113-4
- [43] Castaldi, M.; Ortona, O.; Paduano, L.; Vitagliano, V. *Langmuir* 1998, 14, 5994–5998. doi:10.1021/la980457a
- [44] Camesano, T. A.; Nagarajan, R. *Colloids Surf., A* 2000, 167, 165–177. doi:10.1016/S0927-7757(99)00473-2
- [45] Burilov, V. A.; Mironova, D. A.; Ibragimova, R. R.; Nugmanov, R. I.; Solovieva, S. E.; Antipin, I. S. *Colloids Surf., A* 2017, 515, 41–49. doi:10.1016/j.colsurfa.2016.12.007
- [46] Chakraborty, M.; Panda, A. K. *Spectrochim. Acta, Part A* 2011, 81, 458–465. doi:10.1016/j.saa.2011.06.038
- [47] Gaussian 09, Revision B. 01.; Gaussian, Inc.: Wallingford, CT, 2009.
- [48] Kurzak, B.; Kroczevska, D.; Jezierska, J. *Polyhedron* 1998, 17, 1831–1841. doi:10.1016/S0277-5387(97)00528-7
- [49] Lee, S.; Kim, J.-Y.; Chen, X.; Yoon, J. *Chem. Commun.* 2016, 52, 9178–9196. doi:10.1039/c6cc03584a
- [50] Kolusheva, S.; Zadmard, R.; Schrader, T.; Jelinek, R. J. *Am. Chem. Soc.* 2006, 128, 13592–13598. doi:10.1021/ja064957z
- [51] Thongmalai, W.; Eaidkong, T.; Ampornpun, S.; Mungkarndee, R.; Tumcharern, G.; Sukwattanasinitt, M.; Wacharasindhu, S. J. *Mater. Chem.* 2011, 21, 16391–16397. doi:10.1039/c1jm12795k
- [52] Kolusheva, S.; Kafri, R.; Katz, M.; Jelinek, R. J. *Am. Chem. Soc.* 2001, 123, 417–422. doi:10.1021/ja0034139
- [53] Armarego, W. L. F.; Chai, C. *Purification of Laboratory Chemicals*; Elsevier: New York, 2009; p 743.
- [54] Méndez-Ardoy, A.; Gómez-García, M.; Mellet, C. O.; Sevillano, N.; Girón, M. D.; Salto, R.; Santoyo-González, F.; Fernández, J. M. G. *Org. Biomol. Chem.* 2009, 7, 2681–2684. doi:10.1039/B903635K
- [55] Evans, P. R. *Acta Crystallogr., Sect. D* 2006, 62, 72–82. doi:10.1107/S0907444905036693
- [56] Winn, M. D.; Ballard, C. C.; Cowtan, K. D.; Dodson, E. J.; Emsley, P.; Evans, P. R.; Keegan, R. M.; Krissinel, E. B.; Leslie, A. G. W.; McCoy, A.; McNicholas, S. J.; Murshudov, G. N.; Pannu, N. S.; Potterton, E. A.; Powell, H. R.; Read, R. J.; Vagin, A.; Wilson, K. S. *Acta Crystallogr., Sect. D* 2011, D67, 235–242. doi:10.1107/S0907444910045749
- [57] Battye, T. G. G.; Kontogiannis, L.; Johnson, O.; Powell, H. R.; Leslie, A. G. W. *Acta Crystallogr., Sect. D* 2011, D67, 271–281. doi:10.1107/S0907444910048675
- [58] Sheldrick, G. M. *Acta Crystallogr., Sect. A* 2015, A71, 3–8. doi:10.1107/S2053273314026370
- [59] Sheldrick, G. M. *Acta Crystallogr., Sect. C* 2015, C71, 3–8. doi:10.1107/S2053229614024218
- [60] Stewart, J. J. P. *J. Mol. Model.* 2013, 19, 1–32. doi:10.1007/s00894-012-1667-x
- [61] MOPAC2016; Stewart Computational Chemistry: Colorado Springs, CO, USA, 2016.
- [62] Laikov, D. N.; Ustynyuk, Y. A. *Russ. Chem. Bull.* 2005, 54, 820–826. doi:10.1007/s11172-005-0329-x
- [63] Laikov, D. N. *Chem. Phys. Lett.* 2005, 416, 116–120. doi:10.1016/j.cplett.2005.09.046